

# WEST Search History

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side by side			result set
<i>DB=USPT,PGPB,JPAB,DWPI; PLUR=YES; OP=ADJ</i>			
L2	L1 and IRES and coat protein	2	L2
L1	potato virus X	445	L1

END OF SEARCH HISTORY

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NEWS 7 Apr 22 BIOSIS Gene Names now available in TOXCENTER  
NEWS 8 Apr 22 Federal Research in Progress (FEDRIP) now available  
NEWS 9 Jun 03 New e-mail delivery for search results now available  
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saved answer sets no longer valid  
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now available on STN  
NEWS 20 Aug 19 IFIPAT, IFICDB, and IFIADB have been reloaded  
NEWS 21 Aug 19 The MEDLINE file segment of TOXCENTER has been reloaded  
NEWS 22 Aug 28 Sequence searching in REGISTRY enhanced  
NEWS 23 Sep 03 JAPIO has been reloaded and enhanced  
NEWS 24 Sep 18 Experimental properties added to the REGISTRY file  
NEWS 25 Sep 18 CA Section Thesaurus available in CAPLUS and CA  
NEWS 26 Oct 01 CASREACT Enriched with Reactions from 1907 to 1985  
NEWS 27 Oct 21 EVENTLINE has been reloaded  
NEWS 28 Oct 24 BEILSTEIN adds new search fields  
NEWS 29 Oct 24 Nutraceuticals International (NUTRACEUT) now available on STN  
NEWS 30 Oct 25 MEDLINE SDI run of October 8, 2002  
NEWS 31 Nov 18 DKILUT has been renamed APOLLIT  
NEWS 32 Nov 25 More calculated properties added to REGISTRY  
NEWS 33 Dec 02 TIBKAT will be removed from STN  
NEWS 34 Dec 04 CSA files on STN  
NEWS 35 Dec 17 PCTFULL now covers WP/PCT Applications from 1978 to date  
NEWS 36 Dec 17 TOXCENTER enhanced with additional content  
NEWS 37 Dec 17 Adis Clinical Trials Insight now available on STN  
NEWS 38 Dec 30 ISMEC no longer available  
NEWS 39 Jan 13 Indexing added to some pre-1987 records in CA/CAPLUS  
NEWS 40 Jan 21 NUTRACEUT offering one free connect hour in February 2003  
NEWS 41 Jan 21 PHARMAML offering one free connect hour in February 2003  
NEWS 42 Jan 28 Simultaneous left and right truncation added to COMPENDEX,  
ENERGY, INSPEC

NEWS EXPRESS January 8 CURRENT WINDOWS VERSION IS V8.01a,  
CURRENT MACINTOSH VERSION IS V8.0b(ENG) AND V8.0Jb(JP),  
AND CURRENT DISCOVER FILE IS DATED 01 OCTOBER 2002  
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NEWS PHONE Direct Dial and Telecommunication Network Access to STN  
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\*\*\*\*\* STN Columbus \*\*\*\*\*

FILE 'HOME' ENTERED AT 11:22:42 ON 12 FEB 2003

=> FIL BIOSIS EMBASE CAPLUS  
COST IN U.S. DOLLARS SINCE FILE TOTAL  
ENTRY SESSION  
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FILE 'BIOSIS' ENTERED AT 11:23:27 ON 12 FEB 2003  
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=> s potato virus X  
L1 2178 POTATO VIRUS X

=> s I1 and IRES and coat protein  
L2 4 L1 AND IRES AND COAT PROTEIN

=> dup rem l2  
PROCESSING COMPLETED FOR L2  
L3 2 DUP REM L2 (2 DUPLICATES REMOVED)

=> d bib abs 1-

YOU HAVE REQUESTED DATA FROM 2 ANSWERS - CONTINUE? Y(N):y

L3 ANSWER 1 OF 2 CAPLUS COPYRIGHT 2003 ACS

AN 2002:539852 CAPLUS

DN 137:89449

TI \*\*\*Potato\*\*\* \*\*\*virus\*\*\* \*\*\*X\*\*\* vector containing internal  
ribosomal entry site elements for improved transgene expression in plants  
IN Santa-Cruz, Simon; Pogue, Gregory P.; Toth, Rachel L.; Chapman, Sean;  
Carr, Fiona

PA Biosource Genetics Corporation, USA

SO PCT Int. Appl., 33 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN CNT 1

PATENT NO. KIND DATE APPLICATION NO. DATE

PI WO 2002055719 A2 20020718 WO 2002-US1123 20020109  
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,  
CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,  
GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,  
LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MY, NZ, OM, PH,  
PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,  
UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU,  
TJ, TM  
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH,  
CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, ML, NL, PT, SE, TR,  
BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

PRAI US 2001-758062 A 20010109

AB \*\*\*Potato\*\*\* \*\*\*virus\*\*\* \*\*\*X\*\*\* (PVX)-based vectors were  
generated to investigate use of internal ribosomal entry site (  
\*\*\*IRES\*\*\* ) elements to direct translation of a viral gene. An  
\*\*\*IRES\*\*\* sequence from crucifer-infecting strain of tobacco mosaic  
virus was used to direct expression of the PVX \*\*\*coat\*\*\*  
\*\*\*protein\*\*\*. The \*\*\*IRES\*\*\* was inserted downstream of the gene  
encoding green fluorescent protein (GFP) and upstream of the PVX  
\*\*\*coat\*\*\* \*\*\*protein\*\*\* in either sense or antisense orientation,  
such that \*\*\*coat\*\*\* \*\*\*protein\*\*\* synthesis was dependent on  
ribosome recruitment to the \*\*\*IRES\*\*\*. Stem loop structures were  
inserted at either 3' or 5' end of the \*\*\*IRES\*\*\* to investigate its  
mode of action as these structures block ribosomes. In vitro RNA  
transcripts were inoculated to Nicotiana benthamiana plants and  
protoplasts, levels of GFP and \*\*\*coat\*\*\* \*\*\*protein\*\*\* expression  
were analyzed by ELISA and the rate of viral cell-to-cell movement was  
determined by confocal laser scanning microscopy of GFP synthesis. PVX  
\*\*\*coat\*\*\* \*\*\*protein\*\*\* was expressed, allowing cell-to-cell  
movement of virus, from constructs containing the \*\*\*IRES\*\*\* sequence in  
either sense or antisense orientation and from the construct containing a stem  
loop structure at the 5' end of the \*\*\*IRES\*\*\* sequence. No  
\*\*\*coat\*\*\* \*\*\*protein\*\*\* was synthesized from a construct containing a  
stem loop at the 3' end of the \*\*\*IRES\*\*\* sequence. These results  
suggest that the \*\*\*IRES\*\*\* sequence acts in vivo to direct expression  
of the 3' proximal ORF in a bicistronic mRNA thereby demonstrating the  
potential of employing \*\*\*IRES\*\*\* sequences for the expression of  
foreign proteins from plant virus-based vectors.

L3 ANSWER 2 OF 2 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.DUPLICATE 1

AN 2001:114647 BIOSIS

DN PREV200100114647

TI A novel strategy for the expression of foreign genes from plant virus  
vectors.

AU Toth, Rachel L.; Chapman, Sean; Carr, Fiona; Santa Cruz, Simon (1)  
CS (1) Department of Entomology and Plant Pathology, Horticulture Research  
International, East Malling, Kent, ME19 6BJ; simon.santacruz@hri.ac.uk UK  
SO FEBS Letters, (2 February, 2001) Vol. 489, No. 2-3, pp. 215-219. print.  
ISSN: 0014-5793.

DT Article

LA English

SL English

AB \*\*\*Potato\*\*\* \*\*\*virus\*\*\* \*\*\*X\*\*\* (PVX)-based vector constructs  
were generated to investigate the use of an internal ribosome entry site (  
\*\*\*IRES\*\*\* ) sequence to direct translation of a viral gene. The  
148-nucleotide IREScp sequence from a crucifer-infecting strain of tobacco  
mosaic virus was used to direct expression of the PVX \*\*\*coat\*\*\*  
\*\*\*protein\*\*\* (CP). The \*\*\*IRES\*\*\* was inserted downstream of the  
gene encoding green fluorescent protein (GFP) and upstream of the PVX CP,  
in either sense or antisense orientation, such that CP expression depended  
on ribosome recruitment to the \*\*\*IRES\*\*\*. Stem-loop structures were  
inserted at either the 3'- or 5'-end of the \*\*\*IRES\*\*\* sequence to  
investigate its mode of action. In vitro RNA transcripts were inoculated  
to Nicotiana benthamiana plants and protoplasts: levels of GFP and CP  
expression were analysed by enzyme-linked immunosorbent assay and the rate  
of virus cell-to-cell movement was determined by confocal laser scanning  
microscope imaging of GFP expression. PVX CP was expressed, allowing  
cell-to-cell movement of virus, from constructs containing the  
\*\*\*IRES\*\*\* sequence in either orientation, and from the construct  
containing a stem-loop structure at the 5'-end of the \*\*\*IRES\*\*\*  
sequence. No CP was expressed from a construct containing a stem-loop at  
the 3'-end of the \*\*\*IRES\*\*\* sequence. Our results suggest that the  
\*\*\*IRES\*\*\* sequence is acting in vivo to direct expression of the  
3'-proximal open reading frame in a bicistronic mRNA thereby demonstrating  
the potential of employing \*\*\*IRES\*\*\* sequences for the expression of  
foreign proteins from plant virus-based vectors.

=> s I1 and IRES  
L4 4 L1 AND IRES

=> d hls

(FILE 'HOME' ENTERED AT 11:22:42 ON 12 FEB 2003)

FILE 'BIOSIS, EMBASE, CAPLUS' ENTERED AT 11:23:27 ON 12 FEB 2003  
L1 2178 S POTATO VIRUS X  
L2 4 S L1 AND IRES AND COAT PROTEIN  
L3 2 DUP REM L2 (2 DUPLICATES REMOVED)  
L4 4 S L1 AND IRES

=> s l4 not l2  
L5 0 L4 NOT L2

=> s l1 and vector?

L6 262 L1 AND VECTOR?

=> s16 and coat protein  
L7 66 L6 AND COAT PROTEIN

=> s17 and (chimer? or fusion or heterolog)  
L8 25 L7 AND (CHIMER? OR FUSION OR HETEROLOG)

=> dup rem18  
PROCESSING COMPLETED FOR L8  
L8 16 DUP REM L8 (9 DUPLICATES REMOVED)

=> d16 abs 1-  
YOU HAVE REQUESTED DATA FROM 18 ANSWERS - CONTINUE? Y/(N);y

L9 ANSWER 1 OF 18 CAPLUS COPYRIGHT 2003 ACS  
AN 2002:832988 CAPLUS

DN 137:347521

TI Sequences of synthetic nucleic acid molecule for imparting multiple traits and uses for transforming plants

IN Gonsalves, Dennis; Ferrin-Munoz, Gustavo Alberto

PA Cornell Research Foundation, Inc., USA

SO PCT Int. Appl., 191 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
------------	------	------	-----------------	------

PI	WO 2002086148	A2	20021031	WO 2002-US13377	20020424
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, RW: GH, GM, KE, LS, MW, MZ, SD, SI, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				

PRAI US 2001-286075P P 20010424

AB The present invention is directed to a DNA construct which includes a modified DNA mol. with a nucleotide sequence which is at least 80%, but less than 100%, homologous to two or more desired trait DNA mols. and which imparts the desired trait to plants transformed with the DNA construct. Each of the desired trait DNA mols. relative to the modified nucleic acid mol. have nucleotide sequence similarity values which differ by no more than 3 percentage points. The DNA construct may further include either a silencer or a plurality of modified DNA mols. The present invention also relates to host cells, plant cells, transgenic plants, and transgenic plant seeds contg. such DNA constructs. The present invention is also directed to a method of prep. a modified nucleic acid mol. suitable to impart multiple traits to a plant, a method of detg. whether multiple desired traits can be imparted to plants by a single modified DNA mol., and a method for imparting traits to plants by transforming the plants with the DNA construct.

L9 ANSWER 2 OF 18 CAPLUS COPYRIGHT 2003 ACS

AN 2002:391745 CAPLUS

DN 136:400587

TI DNA vaccines encoding \*\*\*fusion\*\*\* protein of desired antigen and adjuvant sequence of plant viral \*\*\*coat\*\*\* \*\*\*protein\*\*\*

IN Savelyeva, Natalia; Stevenson, Freda

PA Cancer Research Ventures Limited, UK

SO PCT Int. Appl., 84 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
------------	------	------	-----------------	------

PI	WO 2002040513	A2	20020523	WO 2001-GB5142	20011120
WO 2002040513	A3	20021107			
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, RW: GH, GM, KE, LS, MW, MZ, SD, SI, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				

AU 2002023860 A5 20020527 AU 2002-23860 20011120

PRAI GB 2000-28319 A 20001120

WO 2001-GB5142 W 20011120

AB A nucleic acid construct is provided for delivery into living cells in vivo for inducing an immune response in a patient to an antigen; the construct directing the expression of a \*\*\*fusion\*\*\* protein, said \*\*\*fusion\*\*\* protein comprising said antigen and an adjuvant sequence derived from a plant viral \*\*\*coat\*\*\* \*\*\*protein\*\*\*. The plant viral \*\*\*coat\*\*\* \*\*\*protein\*\*\* is \*\*\*potato\*\*\* \*\*\*virus\*\*\*. The antigen is myeloma-specific antigen scFv-5T33, self antigen, tumor antigen, viral antigen derived from e.g. herpes simplex virus or HIV, or bacterial antigen derived from e.g. Staphylococcus or Salmonella. Methods for making such constructs, and methods of using such constructs for the treatment of infectious disease, cancer and B cell malignancy, are provided.

L9 ANSWER 3 OF 18 CAPLUS COPYRIGHT 2003 ACS

AN 2002:10231 CAPLUS

DN 138:84879

TI Production of vaccines using transgenic plants or modified plant viruses as expression \*\*\*vectors\*\*\* and transencapsidated viral coat proteins as epitope presentation systems

IN Hammond, Rosemarie; Zhao, Yan; Hammond, John

PA United States of America, as Represented by the Secretary of Agriculture, USA

SO PCT Int. Appl., 52 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
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PI	WO 2002000169	A2	20020103	WO 2001-US20272	20010828
WO 2002000169	A3	20020718			

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, RW: GH, GM, KE, LS, MW, MZ, SD, SI, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG

AU 2001070164 A5 20020108 AU 2001-70164 20010828

PRAI US 2000-803987 A 20000828

WO 2001-US20272 W 20010828

AB Plants infected with potato virus Y (PVY) were inoculated with infectious PVX RNA encoding PVX \*\*\*coat\*\*\* \*\*\*protein\*\*\* (CP) and modified \*\*\*chimeric\*\*\* NDV/BYMV CP. Antigen-coated plate indirect ELISA and immunoelectron microscopy of virus purified from infected plants showed that progeny virions contained from <1 to as much as 25 \*\*\*chimeric\*\*\* CP. The method can be used to produce and purify large amts. of NDV vaccine. Transencapsidated PVY virions expressing NDV were purified and used to induce anti-NDV antibodies in mice. Further, transgenic tobacco plants expressing HIV/BYMV CP were inoculated with BYMV. Progeny virions from transgenic plants contained transgenic HIV/BYMV CP.

L9 ANSWER 4 OF 18 CAPLUS COPYRIGHT 2003 ACS

AN 2001:195215 CAPLUS

DN 134:232724

TI Plant promoters from the cyclophilin genes of Brassica napus and maize and tomato

IN Gasser, Charles Scott; Budelier, Kim Anne; Gunning, Dorian A.

PA USA

SO U.S., 22 pp.

CODEN: USXXAM

DT Patent

LA English

FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
------------	------	------	-----------------	------

PI	US 6204373	B1	20010320	US 1990-517918	19900502
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PRAI US 1990-517918 19900502

AB The invention provides plant cyclophilin promoters that direct efficient expression of contiguous structural coding sequences in essentially all plant cells and plant organs of transgenic plants. The promoters are isolated using the cDNA sequences encoding cyclophilin from Brassica napus, maize, and tomato. In addn., \*\*\*chimeric\*\*\* genes contg. the plant cyclophilin promoters of the invention and \*\*\*vectors\*\*\* comprising the plant cyclophilin promoters and \*\*\*chimeric\*\*\* genes of the invention are taught herein.

RE.CNT 9 THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 5 OF 18 CAPLUS COPYRIGHT 2003 ACS

AN 2001:573781 CAPLUS

DN 135:271507

TI Plant viral genes in DNA idiotypic vaccines activate linked CD4+ T-cell mediated immunity against B-cell malignancies

AU Savelyeva, Natalia; Munday, Rosalind; Spellerberg, Myfanwy B.;

Lomonosoff, George P.; Stevenson, Freda K.

CS Tenovus Laboratory, Southampton University Hospitals Trust, Southampton,

SO16 6YD, UK

SO Nature Biotechnology (2001), 19(8), 760-764

CODEN: NABIF9, ISSN: 1087-0156

PB Nature America Inc.

DT Journal

LA English

AB DNA delivery of tumor antigens can activate specific immune attack on cancer cells. However, antigens may be weak, and immune capacity can be compromised. \*\*\*Fusion\*\*\* of genes encoding activating sequences to the tumor antigen sequence facilitates promotion and manipulation of effector pathways. Idiotypic determinants of B-cell tumors, encoded by the variable region genes, are clone-specific tumor antigens. When assembled as single-chain Fv (scFv) alone in a DNA vaccine, immunogenicity is low. Previously, the authors found that \*\*\*fusion\*\*\* of a sequence from tetanus toxin (fragment C; FrC) promoted anti-idiotypic protection against lymphoma and myeloma. The authors have now investigated an alternative \*\*\*fusion\*\*\* gene derived from a plant virus, \*\*\*potato\*\*\* \*\*\*virus\*\*\* \*\*\*X\*\*\* \*\*\*coat\*\*\* \*\*\*protein\*\*\*, a primary antigen in humans. When fused to scFv, the self-aggregating protein generates protection against lymphoma and myeloma. In contrast to scFv-FrC, protection against lymphoma is mediated by CD4+ T cells, as is protection against myeloma. Plant viral proteins offer new opportunities to activate immunity against linked T-cell epitopes to attack cancer.

RE.CNT 32 THERE ARE 32 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 6 OF 18 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.DUPLICATE

1

AN 2000:398446 BIOSIS

DN PREV20000398446

TI Transgenic or plant expression \*\*\*vector\*\*\* -mediated recombination of Plum pox virus.

AU Varrelmann, Mark; Palkovics, Laszlo; Maiss, Edgar (1)

CS (1) Institute of Plant Diseases and Plant Protection, University of

Hannover, Herrenhaeuser Str. 2, 30419, Hannover Germany

SO Journal of Virology, (August, 2000) Vol. 74, No. 16, pp. 7482-7489, print. ISSN: 0022-538X.

DT Article

LA English

SL English

AB Different mutants of an infectious full-length clone (p35PPV-NAT) of Plum pox virus (PPV) were constructed: three mutants with mutations of the assembly motifs RQ and DF in the \*\*\*coat\*\*\* \*\*\*protein\*\*\* gene (CP) and two CP \*\*\*chimeras\*\*\* with exchanges in the CP core region of Zucchini yellow mosaic virus and Potato virus Y. The assembly mutants were restricted to single infected cells, whereas the PPV \*\*\*chimeras\*\*\* were able to produce systemic infections in Nicotiana benthamiana plants. After passages in different transgenic N. benthamiana plants expressing the PPV CP gene with a complete (plant line 4.30.45.) or partially deleted 3'-nontranslated region (3'-NTR) (plant line 17.27.4), characterization

of the viral progeny of all mutants revealed restoration of wild-type virus by recombination with the transgenic CP RNA only in the presence of the complete 3'-NTR (4.30.45.). Reconstitution of wild-type virus was also observed following cobombardment of different assembly-defective p35PPV-NAT together with a movement-defective plant expression vector of potato virus X (PVX) expressing the intact PPV-NAT CP gene transiently in nontransgenic *N. benthamiana* plants. Finally, a chimeric recombinant virus was detected after cobombardment of defective p35PPV-NAT with a plant expression vector derived CP gene from the sour cherry isolate of PPV (PPV-SoC). This chimeric virus has been established by a double recombination event between the CP-defective PPV mutant and the intact PPV-SoC CP gene. These results demonstrate that viral sequences can be tested for recombination events without the necessity for producing transgenic plants.

L9 ANSWER 7 OF 16 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.DUPLICATE

AN 2000:278084 BIOSIS  
DN PREV200000278084  
TI Rotavirus VP6 expressed by PVX vectors in *Nicotiana benthamiana* coats PVX rods and also assembles into viruslike particles.  
AU O'Brien, Graham J.; Bryant, Catherine J.; Voogd, Charlotte; Greenberg, Harry B.; Gardner, Richard C.; Bellamy, A. Richard (1)  
CS (1) School of Biological Sciences, University of Auckland, Auckland New Zealand  
SO Virology, (May 10, 2000) Vol. 270, No. 2, pp. 444-453, print.  
ISSN: 0042-6822.  
DT Article  
LA English  
SL English  
AB The rotavirus major inner capsid protein (VP6) has been expressed in *Nicotiana benthamiana* plants using vectors based on potato virus X (PVX). VP6 was expressed either as a fusion with the PVX coat protein or from an additional subgenomic promoter inserted to enable both VP6 and PVX coat protein to be expressed independently. Both approaches yielded VP6, which retained the ability to form trimers. VP6 expressed from the subgenomic promoter assembled into paracrystalline sheets and tubes. Expression as a fusion protein yielded PVX rods that presented an external "overcoat" of VP6, but unexpectedly, some rotavirus protein also assembled into icosahedral viruslike particles (VLPs). The assembly of viral protein into VLPs suggests that prior display of VP6 on the flexuous PVX rod facilitates the subsequent assembly of VP6 into stable icosahedral particles.

L9 ANSWER 8 OF 16 CAPLUS COPYRIGHT 2003 ACS

AN 1998:605027 CAPLUS  
DN 129:198888  
TI DNA construct to confer multiple traits on plants  
IN Pang, Sheng-zhi; Gonsalves, Dennis; Jan, Fuh-yih  
PA Cornell Research Foundation, Inc., USA  
SO PCT Int. Appl., 77 pp.  
CODEN: PIXXD2  
DT Patent  
LA English  
FAN.CNT 1  
PATENT NO. KIND DATE APPLICATION NO. DATE  
PI WO 9837223 A1 19980827 WO 1998-US3030 19980218  
W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, GW, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, UZ, VN, YU, ZW, AM, AZ, BY, BG, BR, BU, CF, CG, CI, CM, FR, GB, GR, IE, IT, LU, MC, ML, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG  
AU 9866571 A1 19980909 AU 1998-66571 19980218  
AU 729308 B2 20010201  
EP 970237 A1 20000112 EP 1998-908568 19980218  
R: CH, DE, FR, GB, IT, LI  
BR 9807587 A 20000321 BR 1998-7587 19980218  
US 2002108146 A1 20020808 US 2001-943215 20010830  
PRAI US 1997-35350 P 19970219  
US 1997-62870 P 19971021  
US 1998-25635 A1 19980218  
WO 1998-US3030 W 19980218  
AB The present invention is directed to a DNA construct formed from a fusion gene which includes a trait DNA mol. and a silencer DNA mol. The trait DNA mol. has a length that is insufficient to impart a desired trait to plants transformed with the trait DNA mol. The silencer DNA mol. is operatively coupled to the trait DNA mol. with the trait and silencer DNA mols. collectively having sufficient length to impart the trait to plants transformed with the DNA construct. Expression systems, host cells, plants, and plant seeds contg. the DNA construct are disclosed. The present invention is also directed to imparting multiple traits to a plant, and in particular to prep. plants which are resistant to multiple viruses. Small nucleocapsid gene fragments (92-235 bp) from tomato spotted wilt virus do not mediate RNA-mediated tospovirus resistance, whereas fusions of the small N gene fragments with jellyfish green fluorescent protein or turnip mosaic virus coat protein are shown to induce RNA-mediated tospovirus resistance.

RE.CNT 9 THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 9 OF 16 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.DUPLICATE

AN 1998:262672 BIOSIS  
DN PREV199800262672  
TI The movement protein of cucumber mosaic virus traffics into sieve elements in minor veins of *Nicotiana glauca*.  
AU Blackman, Leila M.; Boevink, Petra; Cruz, Simon Santa; Palukaitis, Peter; Oparka, Karl J. (1)  
CS (1) Unit Cell Biol., Dep. Virol., Scottish Crop Res. Inst., Invergowrie, Dundee DD2 5DA UK  
SO Plant Cell, (April, 1998) Vol. 10, No. 4, pp. 525-537.  
ISSN: 1040-4651.  
DT Article  
LA English  
AB The location of the 3a movement protein (MP) of cucumber mosaic virus (CMV) was studied by quantitative immunogold labeling of the wild-type 3a

MP in leaves of *Nicotiana glauca* infected by CMV as well as by using a 3a-green fluorescent protein (GFP) fusion expressed from a potato virus X (PVX) vector. Whether expressed from CMV or PVX, the 3a MP targeted plasmodesmata and accumulated in the central cavity of the pore. Within minor veins, the most extensively labeled plasmodesmata were those connecting sieve elements and companion cells. In addition to targeting plasmodesmata, the 3a MP accumulated in the parietal layer of mature sieve elements. Confocal imaging of cells expressing the 3a-GFP fusion protein showed that the 3a MP assembled into elaborate fibrillar formations in the sieve element parietal layer. The ability of 3a-GFP, expressed from PVX rather than CMV, to enter sieve elements demonstrates that neither the CMV RNA nor the CMV coat protein is required for trafficking of the 3a MP into sieve elements. CMV virions were not detected in plasmodesmata from CMV-infected tissue, although large CMV aggregates were often found in the parietal layer of sieve elements and were usually surrounded by 3a MP into sieve elements. CMV virions were not detected in plasmodesmata from CMV-infected tissue, although large CMV aggregates were often found in the parietal layer of sieve elements and were usually surrounded by 3a MP. These data suggest that CMV traffics into minor vein sieve elements as a ribonucleoprotein complex that contains the viral RNA, coat protein, and 3a MP, with subsequent viral assembly occurring in the sieve element parietal layer.

L9 ANSWER 10 OF 16 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.DUPLICATE

AN 1999:74113 BIOSIS  
DN PREV199900074113  
TI Production of a functional single chain antibody attached to the surface of a plant virus.  
AU Smolenska, Lisa (1); Roberts, Ian M.; Learmonth, Deanne; Porter, Andrew J.; Harris, William J.; Wilson, T. Michael A.; Santa Cruz, Simon (1)  
CS (1) Dep. Virol., Scottish Crop Res. Inst., Invergowrie, Dundee DD2 5DA, UK  
SO FEBS Letters, (Dec. 28, 1998) Vol. 441, No. 3, pp. 379-382.  
ISSN: 0014-5793.  
DT Article  
LA English  
AB A potato virus X (PVX) vector was used to express a single chain antibody fragment (scFv) against the herbicide diuron, as a fusion to the viral coat protein. The modified virus accumulated in inoculated *Nicotiana glauca* plants and assembled to give virus particles carrying the antibody fragment. Electron microscopy was used to show that virus particles from infected leaf sap were specifically trapped on grids coated with a diuron-BSA conjugate. The results demonstrate that the PVX vector can be used as a presentation system for functional scFv.

L9 ANSWER 11 OF 16 CAPLUS COPYRIGHT 2003 ACS

AN 1998:191437 CAPLUS  
DN 128:292772  
TI Intracellular location of two groundnut rosette umbravirus proteins delivered by PVX and TMV vectors  
AU Ryabov, E. V.; Oparka, K. J.; Santa Cruz, S.; Robinson, D. J.; Taliansky, M. E.  
CS Virology Dep., Scottish Crop Research Inst., Dundee, DD2 5DA, UK  
SO Virology (1998), 242(2), 303-313  
CODEN: VIRLAX; ISSN: 0042-6822  
PB Academic Press  
DT Journal  
LA English  
AB The proteins encoded by open reading frames (ORF) 3 and 4 of groundnut rosette umbravirus (GRV) were expressed in *Nicotiana benthamiana* as fusions with green fluorescent protein (GFP) from modified potato virus X (PVX) and tobacco mosaic virus (TMV) vectors. Regardless of which plant virus vector was used, GFP fused to the ORF3 protein accumulated in large cytoplasmic inclusion bodies and in nuclei, whereas GFP fused to the ORF4 protein was found in cell walls close to plasmodesmata. Cell-to-cell movement of PVX requires three proteins encoded by the triple gene block (TGB) and also the coat protein (CP). However, when GRV ORF4 was substituted for the PVX CP gene, the hybrid virus was able to move normally in inoculated leaves but not into noninoculated leaves. In contrast, when GRV ORF4 was substituted for the TGB, or for both the TGB and the CP gene, movement of the hybrid viruses was limited to a few epidermal cells neighboring the infection site. Thus, the GRV ORF4 protein can replace the movement proteins of PVX for some of their functions.

L9 ANSWER 12 OF 16 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.DUPLICATE

AN 1998:165030 BIOSIS  
DN PREV199800165030  
TI Simultaneous accumulation of multiple viral coat proteins from a TEV-Nia based expression vector  
AU Ceriani, M. Fernanda; Marcos, Jose F.; Hopp, H. Esteban; Beachy, Roger N. (1)  
CS (1) Dep. Cell Biol., Scripps Res. Inst., 10850 North Torrey Pines Rd., CA 92037 USA  
SO Plant Molecular Biology, (Jan. 2, 1998) Vol. 38, No. 2, pp. 239-248.  
ISSN: 0167-4412.  
DT Article  
LA English  
AB We previously described an expression cassette that relies on the tobacco etch virus (TEV) nuclear inclusion a (Nia) protease and leads to the coordinated accumulation of multiple proteins through self processing of a polyprotein (21). However, low levels of proteins accumulated when the full-length protease was encoded within the polyprotein (22). Studies were conducted to evaluate whether the disruption of Nia nuclear localization would affect the levels of proteins produced via the cassette. Modifications comprised either removal of its nuclear localization signals (NLSs), removal of the VPg domain (which includes the NLSs), and fusion to the 6 kDa protein, previously demonstrated to be a viral cytoplasmic anchor (28). In in vitro translation reactions and in vivo protoplast experiments the modified Nia retained sequence-specific proteolysis. Moreover, the removal of the NLSs correlated with an increase in GUS reporter accumulation. The modified cassette, pPRO10, led to the synthesis of up to three viral coat protein (CPs) in addition to Nia. However, the accumulation of proteins in protoplasts depended upon the position of the CP coding sequence within the cassette as well as on the stability of the protein.

L9 ANSWER 13 OF 16 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.  
AN 1997:309307 BIOSIS  
DN PREV199799617110

TI Restricted virus multiplication in May Queen potato plants transformed with the \*\*\*coat\*\*\* \*\*\*protein\*\*\* gene of potato leafroll Luteovirus.

AU Kondo, Toru (1); Matsumura, Takeshi; Tabayashi, Noriko; Yamashita, Naoko (1); Uyeda, Ichiro (1); Hataya, Tatsuji (1); Saruyama, Haruo; Tanida, Masatoshi; Kimura, Ikuo (1); Shikata, Eishiro

CS (1) Dep. Agrobiol. Bioresources, Fac. Agric., Hokkaido Univ., Kita 9, Nishi 9, Kita-Ku, Sapporo 060 Japan

SO Journal of the Faculty of Agriculture Hokkaido University, (1997) Vol. 67, No. 1, pp. 1-13.

ISSN: 0018-344X.

DT Article

LA English

AB Potato plants (cv. May Queen) transformed with two constructs of the \*\*\*coat\*\*\* \*\*\*protein\*\*\* (CP) gene of potato leafroll virus (PLRV) were produced and analysed for their susceptibility to PLRV. One construct contained only the PLRV CP gene, while the other contained the \*\*\*chimera\*\*\* of \*\*\*potato\*\*\* \*\*\*virus\*\*\* \*\*\*X\*\*\* alpha-beta-leader sequence fused to the PLRV CP gene under the control of the cauliflower mosaic virus 35S promoter. CP transcripts were readily detected by Northern analysis, but CP was not detected by the enzyme-linked immunosorbent assay in transgenic plants. One each from the two constructs of transgenic lines showed restricted virus multiplication at the primary (aphid-borne) infection stage. However, at the secondary (tuber-borne) infection stage, restriction was found to be less effective. Based on these results, the restriction of virus multiplication mediated by CP gene appears to be effective when the inoculum dose is as low as that in a case of aphid inoculation.

L9 ANSWER 14 OF 16 CAPLUS COPYRIGHT 2003 ACS

AN 1996:369872 CAPLUS

DN 125:27694

TI Manufacture of a protein as a \*\*\*fusion\*\*\* product with a viral \*\*\*coat\*\*\* \*\*\*protein\*\*\* with presentation of the protein on the surface of a rod-shaped virus

IN Chapman, Sean Nicholas; Santa Cruz, Simon Peter; Oparka, Karl John; Wilson, Thomas Michael Aubrey

PA Scottish Crop Research Institute, UK

SO PCT Int. Appl., 44 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
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PI WO 9612027	A1	19960425	WO 1995-GB2457	19951018
W:	AM, AT, AU, BB, BG, BR, BY, CA, CH, CN, CZ, DE, DK, EE, ES, FI, GB, GE, HU, IS, JP, KE, KG, KP, KR, KZ, LK, LR, LT, LU, LV, MD, MG, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, TJ, TM, TT			
RW:	KE, MW, SD, SZ, UG, AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG			

CA 2202761	AA	19960425	CA 1995-2202761	19951018
AU 9536598	A1	19960506	AU 1995-36598	19951018
AU 702802	B2	19990304		

EP 787193	A1	19970806	EP 1995-934228	19951018
R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, NL, PT, SE			
US 6232089	B1	20010515	US 1997-844045	19970418

PRAI GB 1994-20680 A 19941018

GB 1995-11729 A 19950609

WO 1995-GB2457 W 19951018

AB A method of manufg. and presenting a protein on the surface of a viral particle is presented. Using a rod-shaped virus the method allows the prodn. of large (i.e. 25 kDa) proteins that assemble with the virus in infected host cells and are arranged on the outer surface of \*\*\*chimeric\*\*\* viruses. The method is particularly intended for use with rod-shaped plant viruses that are relatively benign, such as \*\*\*potato\*\*\* \*\*\*virus\*\*\* \*\*\*X\*\*\*. \*\*\*Potato\*\*\* \*\*\*virus\*\*\* \*\*\*X\*\*\* has the further advantage of being flexuous and so capable of tolerating fairly large inserts in the \*\*\*coat\*\*\* \*\*\*protein\*\*\*. A \*\*\*vector\*\*\* for the prodn. of biol. useful proteins in such a manner is also disclosed. The prodn. of \*\*\*potato\*\*\* \*\*\*virus\*\*\* \*\*\*X\*\*\* presenting green fluorescent protein on its surface and the use of this virus to demonstrate movement of the virus through the plant is described. The \*\*\*fusion\*\*\* protein was linked by a peptide cleavable by foot and mouth disease proteinase 2A and the viral construct included an expression cassette for the proteinase gene. Co-expression of the proteinase gene and the \*\*\*fusion\*\*\* protein gene led to cleavage of some of the \*\*\*fusion\*\*\* protein and generation of a pool of free \*\*\*coat\*\*\* \*\*\*protein\*\*\* to improve the efficiency of formation of the viral coat. Polyprotein processing was essential for the formation of virus capable of systemic propagation through the host plant.

L9 ANSWER 15 OF 16 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.

AN 1996:374740 BIOSIS

DN PREV19969907096

TI Assembly and movement of a plant virus carrying a green fluorescent protein overcoat.

AU Cruz, Simon Santa (1); Chapman, Sean; Roberts, Alison G.; Roberts, Ian M.; Prior, Denton A. M.; Oparka, Karl J.

CS (1) Scottish Crop Res. Inst., Invergowrie, Dundee DD2 9A UK

SO Proceedings of the National Academy of Sciences of the United States of America, (1996) Vol. 93, No. 13, pp. 6286-6290.

ISSN: 0027-8424.

DT Article

LA English

AB \*\*\*Potato\*\*\* \*\*\*virus\*\*\* \*\*\*X\*\*\* (PVX) is a filamentous plant virus infecting many members of the family Solanaceae. A modified form of PVX, PVX.GFP-CP which expressed a \*\*\*chimeric\*\*\* gene encoding a \*\*\*fusion\*\*\* between the 27-kDa Aequorea victoria green fluorescent protein and the amino terminus of the 25-kDa PVX \*\*\*coat\*\*\* \*\*\*protein\*\*\*, assembled into virions and moved both locally and systemically. The PVX.GFP-CP virions were over twice the diameter of wild-type PVX virions. Assembly of PVX.GFP-CP virions required the presence of free \*\*\*coat\*\*\* \*\*\*protein\*\*\* subunits in addition to the \*\*\*fusion\*\*\* protein subunits. PVX.GFP-CP virions accumulated as paracrystalline arrays in infected cells similar to those seen in cells

infected with wild-type PVX. The formation of virions carrying large superficial fusions illustrates a novel approach for production of high levels of foreign proteins in plants. Aggregates of PVX.GFP-CP particles were fluorescent, emitting green light when excited with ultraviolet light and could be imaged using confocal laser scanning microscopy. The detection of virus particles in infected tissue demonstrates the potential of fusions between the green fluorescent protein and virus \*\*\*coat\*\*\* \*\*\*protein\*\*\* for the non-invasive study of virus multiplication and spread.

L9 ANSWER 16 OF 16 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.DUPLICATE

AN 1996:123580 BIOSIS

DN PREV199698695725

TI Imaging the green fluorescent protein in plants-viruses carry the torch.

AU Oparka, K. J. (1); Roberts, A. G.; Prior, D. A. M.; Chapman, S.; Baulcombe, D.; Santa Cruz, S.

CS (1) Scottish Crop Research Inst., Invergowrie, Dundee DD2 5DA UK

SO Protoplasma, (1995) Vol. 189, No. 3-4, pp. 133-141.

ISSN: 0033-183X.

DT General Review

LA English

AB The green fluorescent protein (GFP) was introduced into plant cells using \*\*\*potato\*\*\* \*\*\*virus\*\*\* \*\*\*X\*\*\* as a \*\*\*vector\*\*\*. The GFP was produced at high levels within virus-infected cells by utilising a duplication of the viral \*\*\*coat\*\*\* \*\*\*protein\*\*\* subgenomic RNA promoter sequence to direct transcription of mRNA encoding the GFP. We also exploited the ability of GFP to retain its fluorescence when fused to other proteins by fusing it to the PVX \*\*\*coat\*\*\* \*\*\*protein\*\*\*. The resultant fluorescent virus became systemic and its movement from cell to cell was traced using confocal laser scanning microscopy. Using PVX as the \*\*\*vector\*\*\*, additional fusions of the GFP were made to the movement protein of tobacco mosaic virus (TMV). The fluorescent \*\*\*fusion\*\*\* protein produced was targeted to specific wall sites thought to be plasmodesmata pit fields. The utility of virus-based \*\*\*vectors\*\*\* for the delivery and targeting of GFP in living plant cells is discussed.

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FULL ESTIMATED COST		98.57	98.78

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)	ENTRY	SINCE FILE	TOTAL
CA SUBSCRIBER PRICE		-5.86	-5.88

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PASSWORD:

\*\*\*\*\* RECONNECTED TO STN INTERNATIONAL \*\*\*\*\*

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FILE 'HOME' ENTERED AT 17:18:18 ON 20 FEB 2003

COST IN U.S. DOLLARS	ENTRY	SINCE FILE	TOTAL
FULL ESTIMATED COST		0.21	0.21

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COST IN U.S. DOLLARS		0.21	0.21
FULL ESTIMATED COST		0.21	0.21

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=> s IRES and viral vector  
L1 40 IRES AND VIRAL VECTOR

=> dup rem l1  
PROCESSING COMPLETED FOR L1  
L2 29 DUP REM L1 (11 DUPLICATES REMOVED)

=> s l2 and py<2001  
2 FILES SEARCHED...  
L3 14 L2 AND PY<2001

=> d bib abs 1-  
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L3 ANSWER 1 OF 14 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.  
AN 2000:178123 BIOSIS

DN PREV200000179123

TI Establishment of efficient reaggregation culture system for gene transfection into immature T cells by retroviral vectors.

AU Hozumi, Katsuto; Ohtsuka, Ryo; Suzuki, Daisuke; Ando, Kiyoshi; Ito, Mamoru; Nishimura, Takashi; Merkensthaler, Matthias; Habu, Sonoko (1)  
CS (1) Department of Immunology, Tokai University School of Medicine, Bohseidai, Isehara, 259-1193 Japan  
SO Immunology Letters, ( \*\*\*Jan. 10, 2000\*\*\* ) Vol. 71, No. 1, pp. 81-86.  
ISSN: 0165-2478.  
DT Article  
LA English  
SL English

AB To overcome low efficiency of retroviral infection into immature T cells, we modified reaggregation fetal thymus organ culture by closely packed co-culture with virus-producing cells (VPC). The \*\*\*viral\*\*\* vector was constructed in chimeric vector, pMX, with \*\*\*IRES\*\*\* and tailless-rat CD2 as a surface marker of infected cells. A rearranged TCR beta gene (Vbeta8.2) was further inserted into the construct for investigating effect of the introduced gene in T cell development. Using this system, we succeeded to transfer the \*\*\*viral\*\*\* vector into immature thymocytes at a remarkably higher efficiency compared to conventional methods using medium containing retrovirus. Moreover, the introduced TCR beta gene was expressed on thymocytes of RAG2-deficient mice to induce in the transition of CD4-CD8- double-negative (DN) into CD4+CD8+ double-positive (DP) cells by transducing beta-selection signaling. Thus, our modified reaggregation culture system is useful for studying the molecular mechanism of T cell development due to a highly efficient gene transfer into immature T cells.

L3 ANSWER 2 OF 14 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.  
AN 2000:68467 BIOSIS  
DN PREV200000086467

TI Prevention of 6-hydroxydopamine-induced rotational behavior by BDNF somatic gene transfer.

AU Klein, Ronald L. (1); Lewis, Mark H.; Muzyczka, Nicholas; Meyer, Edwin M.  
CS (1) Department of Pharmacology and Therapeutics, University of Florida, Gainesville, FL USA

SO Brain Research, ( \*\*\*Nov. 20, 1999\*\*\* ) Vol. 847, No. 2, pp. 314-320.  
ISSN: 0006-8993.

DT Article  
LA English  
SL English

AB Brain-derived neurotrophic factor (BDNF) was expressed via injection of \*\*\*viral\*\*\* vector into the substantia nigra pars compacta (SNc) to investigate its influence on nigrostriatal dopaminergic activity and locomotor behavior. The recombinant adeno-associated virus (AAV) vector, pTR-BDNFmyc, incorporated the neuron-specific enolase (NSE) promoter and the internal ribosome entry site (IRES) element driving expression of both epitope-tagged BDNF and green fluorescent protein (GFP) bidirectionally. The control vector, pTR-UF4, incorporated NSE promoter-driven GFP expression only. Transgene expression persisted in both vector groups throughout the 9 month course of the study. Partial 6-hydroxydopamine (6-OHDA) lesions were conducted in the SNc ipsilateral to, and 6 months after, transduction with either the pTR-BDNFmyc or the pTR-UF4. Transgenic BDNFmyc had no effect on the number of tyrosine hydroxylase (TH)-labeled neurons in the SNc after 6-OHDA-lesions, but did block the amphetamine-induced, ipsiversive, turning-behavior caused by the lesion in the pTR-UF4 group. The BDNFmyc-transduced group also demonstrated more locomotor activity and rotational activity contralateral to the lesioned side than did the pTR-UF4-transduced group. Long-term, stable expression of BDNF can therefore modulate locomotor activity without significantly affecting nigrostriatal dopaminergic survival.

L3 ANSWER 3 OF 14 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.  
AN 1999:126783 BIOSIS  
DN PREV199900126783

TI Antisense oligonucleotide inhibition of hepatitis C virus (HCV) gene expression in livers of mice infected with an HCV-vaccinia virus recombinant.

AU Zhang, Hong; Hanecak, Ronnie; Brown-Driver, Vickie; Azad, Raana; Conklin, Boyd; Fox, Maureen C.; Anderson, Kevin P. (1)

CS (1) 2292 Faraday Ave., Carlsbad, CA 92008 USA

SO Antimicrobial Agents and Chemotherapy, ( \*\*\*Feb., 1999\*\*\* ) Vol. 43, No. 2, pp. 347-353.  
ISSN: 0066-4804.

DT Article  
LA English

AB Hepatitis C virus (HCV) is the major cause of non-A, non-B hepatitis worldwide. Current treatments are not curative for most infected individuals, and there is an urgent need for both novel therapeutic agents and small-animal models which can be used to evaluate candidate drugs. A small-animal model of HCV gene expression was developed with recombinant vaccinia virus vectors. VHCV-\*\*\*IRES\*\*\* (internal ribosome entry site) is a recombinant vaccinia \*\*\*viral\*\*\* vector containing the HCV 5' nontranslated region (5'-NTR) and a portion of the HCV core coding region fused to the firefly luciferase gene. Intraperitoneal injection of VHCV-\*\*\*IRES\*\*\* produced high levels of luciferase activity in the livers of BALB/c mice. Antisense oligonucleotides complementary to the HCV 5'-NTR and translation initiation codon regions were then evaluated for their effects on the expression of these target HCV sequences in BALB/c mice infected with the vaccinia virus vector. Treatment of VHCV-\*\*\*IRES\*\*\*-infected mice with 20-base phosphorothioate oligonucleotides complementary to the sequence surrounding the HCV initiation codon (nucleotides 330 to 349) specifically reduced luciferase expression in the livers in a dose-dependent manner. Inhibition of HCV reporter gene expression in this small-animal model suggests that antisense oligonucleotides may provide a novel therapy for treatment of chronic HCV infection.

L3 ANSWER 4 OF 14 CAPLUS COPYRIGHT 2003 ACS  
AN 2001:480838 CAPLUS  
DN 135:87978

TI Mammalian retroviral vectors and their uses in study of gene expression

IN Beach, David H.; Hannon, Gregory J.; Conklin, Douglas; Sun, Peiqing  
PA Cold Spring Harbor Laboratory, USA

SO U.S., 80 pp., Cont.-in-part of U.S. 6,025,192.  
CODEN: USXXAM

DT Patent  
LA English

FAN.CNT 2

PATENT NO.	KIND DATE	APPLICATION NO. DATE
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PI US 6256071	B1 20010703	US 1997-820831 19970319
US 6025192	A 20000215	US 1996-718928 19960920 <-
CA 2262476	AA 19980326	CA 1997-226276 19970922 <-

WO 9812339 A2 19980326 WO 1997-US17579 19970922 <-  
WO 9812339 A3 19980903

W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, HU, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM  
RW: GH, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG

AU 9746590 A1 19980414 AU 1997-46590 19970922 <-  
AU 738156 B2 20010913

EP 932695 A2 19990804 EP 1997-945369 19970922 <-  
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI

JP 2002514054 T2 20020514 JP 1998-515028 19970922

PRAI US 1996-718928 A2 19960920

US 1997-820931 A 19970319

WO 1997-US17579 W 19970922

AB The present invention relates to methods and comps. for the elucidation of mammalian gene function. Expression vectors for animal cells that use regulatory elements of retroviruses to drive expression of cloned genes are described. These vectors are replication-defective and can be used in improved mammalian complementation screening, functional inactivation of specific essential or non-essential mammalian genes, and identification of mammalian genes modulated by specific stimuli. Construction of plasmids for the manuf. of a no. of such vectors is described. In particular, the comps. of the present invention include, but are not limited to, replication-deficient retroviral vectors, libraries comprising such vectors, retroviral particles produced by such vectors in conjunction with retroviral packaging cell lines, integrated provirus sequences derived from the retroviral particles of the invention and circularized provirus sequences which have been excised from the integrated provirus sequences of the invention. The comps. of the present invention further include novel retroviral packaging cell lines.

RE.CNT 63 THERE ARE 63 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 5 OF 14 CAPLUS COPYRIGHT 2003 ACS

AN 2000:881527 CAPLUS

DN 134:28054

TI A novel packaging cell line for the rescue, production and titration of high-capacity adenovirus vectors

IN Krouglik, Valeri A.; Eisenmith, Randy C.

PA Mount Sinai School of Medicine, USA

SO PCT Int. Appl., 53 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

PATENT NO.	KIND DATE	APPLICATION NO. DATE
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PI WO 2000072887 A1 20001207 WO 2000-US14914 20000526 <-  
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, MY, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM  
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG

PRAI US 1999-136481 P 19990528

AB The present invention describes a method of producing adenovirus gutless amplicon \*\*\*viral\*\*\* vector substantially reduced in the content of helper virus. The invention also describes a system for the helper virus independent replication and packaging of adenovirus gutless vectors. The method avoids the problem by placing helper functions on an episome based on an Epstein-Barr virus replicon that is stable at a low copy no but that lacks the encapsidation signal and the terminal protein gene. The helper functions are under control of a regulated promoter. Viral replication is induced when the cells are transformed with a vector carrying the terminal protein gene. A cell line for this system is also discussed.

RE.CNT 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 6 OF 14 CAPLUS COPYRIGHT 2003 ACS

AN 2000:209936 CAPLUS

DN 132:246355

TI Methods using beta.-endorphin-expressing recombinant expression systems for treating chronic pain

IN Iadarola, Michael J.; Caudle, Robert M.; Finegold, Alan A.; Mannes, Andrew J.; Olah, Zoltan

PA Government of the United States of America, Represented by the Secretary, Department of Health and Human Services, USA

SO PCT Int. Appl., 49 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 2

PATENT NO.	KIND DATE	APPLICATION NO. DATE
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PI WO 2000018800 A2 20000330 WO 1999-US22103 19990923 <-  
WO 2000018800 A3 20000720

W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, GR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM  
RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG

AU 9962809 A1 20000410 AU 1999-62809 19990923 <-

PRAI US 1998-100910 P 19980923

WO 1999-US22103 W 19990923

AB Comps. and methods are provided which selectively treat chronic pain while not significantly affecting basal nociceptive, acute pain responses. The invention provides for comps. and methods of treating chronic pain by administering beta.-endorphin-expressing recombinant expression systems, e.g. adenovirus or adeno-assoc. virus, into a subarachnoid or epidural

space. The recombinant virus infects the pia mater connective tissue cells and the infected cells express the fusion protein, wherein the fusion protein is secreted into the spinal cord parenchymal tissue in an amt. effective to treat the chronic pain but not significantly affecting basal nociceptive responses.

L3 ANSWER 7 OF 14 CAPLUS COPYRIGHT 2003 ACS

AN 2000:20 CAPLUS

DN 132:32667

TI Cloning of Escherichia coli cytosine deaminase gene and expression of the gene using a new \*\*\*viral\*\*\* \*\*\*vector\*\*\*

IN Gu, Jianren; Ren, Shengjun; Xu, Xulan

PA Shanghai Tumour Research Institute, Peop. Rep. China

SO Faming Zhuanti Shengding Gongkai Shuomingshu, 39 pp.

CODEN: CNXXEV

DT Patent

LA Chinese

FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
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PI	CN	1161375	A	19971008	CN	1998-116598	19981129	<-
	CN	1055968	B	20000830				

PRAI CN 1998-116598 19981129

AB The gene encoding cytosine deaminase of Escherichia coli strain H-30 was cloned, and its initiation codon of 'GTG' was mutated to 'ATG' by PCR. Prepn. of prokaryotic recombinant expression vector pBV220-CD; prepn. of packaging cells for producing infectious pseudo-retrovirus or pseudo-adenovirus vectors; and use of the pseudo-virus for treating cancer along with 5-FC (5-fluorocytosine), which induces lethal toxicity to the cells contg. active CD gene, are also described.

L3 ANSWER 8 OF 14 CAPLUS COPYRIGHT 2003 ACS

AN 1999:436294 CAPLUS

DN 131:69280

TI Novel gene trap and its use for high efficiency selection of regulated eukaryotic genes

IN Baetscher, Manfred; Nir, Waan-jeng

PA Biotransplant, Inc., USA

SO U.S., 23 pp., Cont. of U.S. Ser. No. 374,833, abandoned.

CODEN: USXXAM

DT Patent

LA English

FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
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PI	US	5922601	A	19990713	US	1996-716854	19980916	<-
	PRAI	US	1995-374833	19950119				

AB The invention provides a novel gene trap construct that allows for high efficiency identification and selection of eukaryotic genes whose activity is regulated upon a cellular transition. Said \*\*\*viral\*\*\* \*\*\*vector\*\*\* comprises in its downstream sequence (i) a cassette having a functional splice acceptor, a translation stop sequence and an internal ribosome entry site and (ii) a promoterless protein coding sequence encoding at least one polypeptide providing pos. and neg. selection traits. Also provided is a method for identification of genes whose activity is regulated upon a cellular transition event by introducing the gene trap construct into a cell and observing expression of the pos. and/or neg. selection traits before and after the transition event.

RE.CNT 21 THERE ARE 21 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 9 OF 14 CAPLUS COPYRIGHT 2003 ACS

AN 1999:194259 CAPLUS

DN 130:23258

TI \*\*\*viral\*\*\* \*\*\*vector\*\*\* system capable of expressing an apoptosis-associated gene

IN Hamada, Hirochika

PA RPR Gencell Asia/Pacific Inc., Japan

SO PCT Int. Appl., 51 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
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PI	WO	9913073	A2	19990318	WO	1998-JP4010	19980907	<-
	WO	9913073	A3	19990610				

W: AU, CA, KR, NZ, US

RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE

JP 11075859 A2 19990323 JP 1997-259235 19970908 <-

AU 8889991 A1 19990329 AU 1998-89991 19980907 <-

PRAI JP 1997-259235 19970908

WO 1998-JP4010 19980907

AB An apoptosis-resistant virus-sensitive cell line based upon cell line 293 is disclosed. To such cells, apoptosis resistance genes such as crmA, bcl-2, bcl-x1, FLIP, survivin, IAP, or ILP have been introduced. The generation of adenovirus vectors capable of expressing apoptosis-assoc. genes such as FAS, FLICE, bcl-x2, and Bax is achieved using said cell line. The recombinant viruses of the invention may be useful for gene therapy for cancer, autoimmune diseases, graft rejection, and inflammatory diseases.

L3 ANSWER 10 OF 14 CAPLUS COPYRIGHT 2003 ACS

AN 1998:395891 CAPLUS

DN 129:131842

TI In vivo expression of therapeutic human genes for dopamine production in the caudates of MPTP-treated monkeys using an AAV vector

AU Duling, M. J.; Samulski, R. J.; Elsworth, J. D.; Kaplitt, M. G.; Leone, P.; Xiao, X.; Li, J.; Freese, A.; Taylor, J. R.; Roth, R. H.; Sladek, J. R., Jr.; O'malley, K. L.; Redmond, D. E., Jr.

CS Department of Molecular Medicine, University of Auckland School of Medicine, Auckland, N. Z.

SO Gene Therapy ( 1998 ), 5(6), 820-827

CODEN: GETHEC; ISSN: 0899-7128

PB Stockton Press

DT Journal

LA English

AB An adeno-assoc. virus (AAV) vector, expressing genes for human tyrosine hydroxylase (TH) and arom. amino acid decarboxylase (AADC), demonstrated significantly increased prodn. of dopamine in 293 (human embryonic kidney) cells. This biostronic vector was used to transduce striatal cells of

six asymptomatic but dopamine-depleted monkeys which had been treated with the neurotoxin MPTP. Striatal cells were immunoreactive for the vector-encoded TH after stereotactic injection for periods up to 134 days, with biochem. effects consistent with dopamine biosynthetic enzyme expression. A subsequent expt. was carried out in six more severely depleted and parkinsonian monkeys. Several TH-aadc-treated monkeys showed elevated levels of dopamine near injection tracts after 2.5 mo. Two monkeys that received a .beta.-galactosidase expressing vector showed no change in striatal dopamine. Behavioral changes could not be statistically related to the vector treatment groups. Toxicity was limited to transient fever in several animals and severe hyperactivity in one animal in the first days after injection with no assoc. histol. evidence of inflammation. This study shows the successful transfection of primate neurons over a period up to 2.5 mo with suggestive evidence of biochem. phenotypic effects and without significant toxicity. While supporting the idea of an in vivo gene therapy for Parkinson's disease, more consistent and longer lasting biochem. and behavioral effects will be necessary to establish the feasibility of this approach in a primate model of parkinsonism.

RE.CNT 28 THERE ARE 28 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 11 OF 14 CAPLUS COPYRIGHT 2003 ACS

AN 1998:89371 CAPLUS

DN 128:150403

TI Construction of retroviral vectors for delivering viral and oncogenic inhibitors

IN Raybak, Susanna M.; Cara, Andrea; Gusella, Gabriele Luca; Newton, Dianne L.

PA United States Dept. of Health and Human Services, USA; Raybak, Susanna M.; Cara, Andrea; Gusella, Gabriele Luca; Newton, Dianne L.

SO PCT Int. Appl., 63 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
------------	------	------	-----------------	------

PI	WO	9803669	A2	19980129	WO	1997-US12637	19970717	<-
	WO	9803669	A3	19980228				

W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, HU, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM

RW: GH, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG

AU 9738049 A1 19980210 AU 1997-38049 19970717 <-

AU 734968 B2 20010828

EP 917585 A2 19990528 EP 1997-935014 19970717 <-

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI

PRAI US 1996-22052P P 19960722

WO 1997-US12637 W 19970717

AB Cell transformation vectors for inhibiting HIV and tumor growth are provided. Optionally, the vectors encode RNases A superfamily members such as eosinophil-derived neurotoxin (EDN) and onconase. Cells transduced by the vectors and methods of transforming cells (in vitro and in vivo) using the vectors are also provided. The viral and oncogene inhibitors are typically linked to a promoter such as retroviral HIV LTR promoters, the CMV promoter, the probasin promoter, and tetracycline-responsive promoters. The method is exemplified by construction of a \*\*\*viral\*\*\* \*\*\*vector\*\*\* contg. a HIV Rev-responsive element, an encephalomyocarditis virus internal ribosome entry site, a first viral inhibitor subsequence (for immunodominant proteins such as Tat, Gag, or Rev), splice donor site subsequence, splice acceptor site subsequence, the above mentioned promoter, and the EDN coding sequence. The vector may be packaged in a liposome and its contents transduced into CD34+ hematopoietic stem cells, CD4+ cells, and transferin receptor+ cells. Claimed vectors include pBAR, pBAR-ONC, and pBAR-EDN.

L3 ANSWER 12 OF 14 CAPLUS COPYRIGHT 2003 ACS

AN 1998:15859 CAPLUS

DN 128:85136

TI Construction of adenoviral gene vectors for mammalian cells

IN Perricaudet, Michel; Yeh, Patrice; Leblois-Prehaud, Helene

PA Rhone-Poulenc Rorer S.A., Fr.; Perricaudet, Michel; Yeh, Patrice; Leblois-Prehaud, Helene

SO PCT Int. Appl., 55 pp.

CODEN: PIXXD2

DT Patent

LA French

FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
------------	------	------	-----------------	------

PI	WO	9747757	A1	19971218	WO	1997-FR914	19970523	<-
	W:	AL, AU, BA, BB, BG, BR, CA, CN, CU, CZ, EE, GE, GH, HU, IL, IS, JP, KP, KR, LC, LK, LR, LT, LV, MG, MK, MN, MX, NO, NZ, PL, RO, SG, SI, SK, TR, TT, UA, US, UZ, VN, YU, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM						

RW: GH, KE, LS, MW, SD, SZ, UG, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG

FR 2749857 A1 19971219 FR 1996-7273 19960812 <-

FR 2749857 B1 19980814

CA 2257916 AA 19971218 CA 1997-2257916 19970523 <-

AU 9730377 A1 19980107 AU 1997-30377 19970523 <-

AU 726442 B2 20001109

EP 906443 A1 19990407 EP 1997-925133 19970523 <-

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, PT, IE, SI, FI

BR 9709700 A 19990810 BR 1997-9700 19970523 <-

JP 2000511779 T2 20000912 JP 1998-501269 19970523 <-

NO 9805739 A 19981208 NO 1998-5739 19981208 <-

KR 2000016524 A 20000325 KR 1998-710112 19981210 <-

PRAI FR 1996-7273 A 19960812

WO 1997-FR914 W 19970523

AB The invention discloses circular and replicating DNA mols., useful in gene therapy, as well as a particularly efficient method for generating them in situ from a mutant adenovirus-derived vector. The adenovirus carries a

deletion mutation in the E1 gene. The DNA sequences carried by the adenoviral vectors are a gene of interest, replication origins from viruses such as the Epstein-Barr virus (EBV) and papillomavirus, ARS sequences, and an inducible promoter controlling the Cre recombinase gene. The promoter is derived from mouse mammary tumor virus and is inducible by dexamethason or tetracycline. The viral replication origin regions are dependent on site-specific recombination. The viral vectors also contain inverted repeat sequences from the P1 phage *loxP* region which are responsive to Cre recombinase. The method is exemplified by constructing a \*\*\*viral\*\*\* \*\*\*vector\*\*\* contg. the EBV EBNA1 gene and *oriP* region, a mammalian cell-functional gene promoter, and the \*\*\*IRES\*\*\* genetic element from encephalomyocarditis virus.

L3 ANSWER 13 OF 14 CAPLUS COPYRIGHT 2003 ACS

AN 1998:567287 CAPLUS

DN 125:187592

TI RNA virus vector and helper virus or cell line for gene cloning, vaccine development, and neoplasm and inflammation inhibitor recombinant production

IN Mertelsmann, Roland; Rosenthal, Felicia; Kalden, Joachim; Bertling, Wolf; Lindemann, Albrecht; Kulmburg, Peter; Veelken, Hendrik

PA Klinikum der Albert-Ludwigs-Universitaet Freiburg, Germany

SO Ger. Offen., 11 pp.

CODEN: GWXXBX

DT Patent

LA German

FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI DE 19503082	A1	19960808	DE 1995-19503082	19950201 <-
WO 9623689	A1	19960808	WO 1996-EP334	19960129 <-
W:	AU, BR, CA, CN, JP, KR, RU, US			
RW:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE			
AU 9652598	A1	19960821	AU 1996-52598	19960129 <-
EP 804588	A1	19971105	EP 1996-900980	19960129 <-
R:	AT, BE, CH, DE, FR, GB, IT, LI, NL			
US 6255104	B1	20010703	US 1998-894170	19980512

PRAI DE 1995-19503082 A 19950201

WO 1996-EP334 W 19960129

AB RNA virus vectors in conjunction with helper viruses or helper cell lines are useful for gene cloning. Recombinant neoplasm inhibitors and inflammation inhibitors can be produced by this method. Vaccine development is another application. An example is poliovirus interleukin-2 gene expression in tumor treatment. The EMC \*\*\*IRES\*\*\* element was used in the poliovirus vector. Another example is human gene fas expression using a \*\*\*viral\*\*\* \*\*\*vector\*\*\* to induce cell type-specific apoptosis.

L3 ANSWER 14 OF 14 CAPLUS COPYRIGHT 2003 ACS

AN 1993:206917 CAPLUS

DN 118:206917

TI Characterization of a bicistronic retroviral vector composed of the swine vesicular disease virus internal ribosome entry site

AU Chen, Bing Fang; Hwang, Lih Hwa; Chen, Ding Shinn

CS Coll. Med., Natl. Taiwan Univ., Taipei, Taiwan

SO Journal of Virology ( \*\*\*1993\*\*\* ), 67(4), 2142-8

CODEN: JOVIAH; ISSN: 0022-538X

DT Journal

LA English

AB The 5' nontranslated region (NTR) from the genome of swine vesicular disease virus (SVDV), a member of the family Picornaviridae, was cloned and used to construct a bicistronic retroviral vector. The vector is characterized by coexpression of 2 genes from a single transcript. Inclusion of the 5' NTR of SVDV did not negate the \*\*\*viral\*\*\* \*\*\*vector\*\*\*. Protein anal. indicated that the 5' NTR could efficiently direct internal initiation, thus allowing the downstream gene to be translated. Translation of the internally initiated porcine growth hormone gene was approx. 30-fold less than that when the porcine growth hormone gene was at the upstream position in NIH 3T3 cells but was about equiv. to that in HeLa cells, implying that some cellular factors that stimulated internal initiation of the SVDV 5' NTR are present in HeLa cells. However, in G418-selected clones, the NeoR-encoding gene was expressed with equiv. efficiency either at a downstream position or at an upstream position in either NIH 3T3 or HeLa cells. Compared with the conventional double-gene vector or the U3-based vector, the bicistronic vector coexpressed 2 genes much more efficiently, owing to elimination of promoter interference. Furthermore, this type of vector infected and expressed the target genes efficiently in 2 primary cell lines, rat embryo and human skin fibroblast cells, that were tested. These expl. data suggest a better design for the retroviral vector and provide evidence that internal initiation of the SVDV 5' NTR was stimulated cell-specifically.

=> d his

(FILE 'HOME' ENTERED AT 17:17:50 ON 20 FEB 2003)

FILE 'BIOSIS, EMBASE, CAPLUS' ENTERED AT 17:18:32 ON 20 FEB 2003

L1 40 S IRES AND VIRAL VECTOR

L2 28 DUP REM L1 (11 DUPLICATES REMOVED)

L3 14 S L2 AND PY<2001

=> s IRES and retrovir?

L4 505 IRES AND RETROVIR?

=> s 14 and py<2001

1 FILES SEARCHED...

L5 324 L4 AND PY<2001

=> d bib abs 1-10

L5 ANSWER 1 OF 324 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.

AN 2002:425400 BIOSIS

DN PREV200200425400

TI Rous sarcoma virus translation revisited: Characterization of an internal ribosome entry segment in the 5' leader of the genomic RNA.

AU Deffaud, Clarence; Darlix, Jean-Luc (1)

CS (1) Laboratoire, Unite de Virologie Humaine, Institut National de la Sante et de la Recherche Medicale, Ecole Normale Supérieure de Lyon, 46 Allée d'Italie, No. 412, 69364, Lyon Cedex 07; Jean-Luc.Darlix@ens-lyon.fr France

SO Journal of Virology, ( \*\*\*December, 2000\*\*\* ) Vol. 74, No. 24, pp.

11581-11588. <http://intl-vi.asm.org/>. print.

ISSN: 0022-538X.

DT Article

LA English

AB The 5' leader of Rous sarcoma virus (RSV) genomic RNA and of \*\*\*retroviruses\*\*\* in general is long and contains stable secondary structures that are critical in the early and late steps of virus replication such as RNA dimerization and packaging and in the process of reverse transcription. The initiation of RSV Gag translation has been reported to be 5' cap dependent and controlled by three short open reading frames located in the 380-nucleotide leader upstream of the Gag start codon. Translation of RSV Gag would thus differ from that prevailing in other \*\*\*retroviruses\*\*\* such as murine leukemia virus, reticuloendotheliosis virus type A, and simian immunodeficiency virus, in which an internal ribosome entry segment ( \*\*\*IRES\*\*\* ) in the 5' end of the genomic RNA directs efficient Gag expression despite stable 5' secondary structures. This prompted us to investigate whether RSV Gag translation might be controlled by an \*\*\*IRES\*\*\* -dependent mechanism. The results show that the 5' leaders of RSV and v-Src RNA exhibit \*\*\*IRES\*\*\* properties, since these viral elements can promote efficient translation of monocistronic RNAs in conditions inhibiting 5' cap-dependent translation. When inserted between two cistrons in a canonical bicistronic construct, both the RSV and v-Src leaders promote expression of the 3' cistron. A genetic analysis of the RSV leader allowed the identification of two nonoverlapping 5' and 3' leader domains with \*\*\*IRES\*\*\* activity. In addition, the v-Src leader was found to contain unique 3' sequences promoting an efficient reinitiation of translation. Taken together, these data lead us to propose a new model for RSV translation.

L5 ANSWER 2 OF 324 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.

AN 2001:322188 BIOSIS

DN PREV200100322188

TI Restoration of WASP-deficient T-cell signaling defects in mice upon transplantation of \*\*\*retrovirally\*\*\* transduced hematopoietic stem cells.

AU Klein, Christoph (1); Nguyen, Deanna; Liu, Ching-Hui; Rosen, Fred S.; Alt, Fred W.; Mulligan, Richard C.; Snapper, Scott B.

CS (1) Pediatric Hematology/Oncology, Medical School Hannover, Hannover Germany

SO Blood, ( \*\*\*November 16, 2000\*\*\* ) Vol. 96, No. 11 Part 1, pp. 591a.

print.

Meeting Info.: 42nd Annual Meeting of the American Society of Hematology San Francisco, California, USA December 01-05, 2000 American Society of Hematology

. ISSN: 0006-4971.

DT Conference

LA English

SL English

AB Wiskott Aldrich Syndrome, an X-linked primary immunodeficiency disorder, is caused by mutations in the WASP gene. WASP plays a critical role in antigen-receptor signaling in T-cells by regulating cell surface signals to the cytoskeleton. In both patients and in WASP KO mice, WASP-deficient T cells do not proliferate and have cytoskeletal defects in response to CD3 crosslinking. In this study, we have used a WASP knockout mouse model to assess whether transplantation of \*\*\*retrovirally\*\*\* transduced hematopoietic stem cells would reverse the specific T-cell signaling deficiency. First, we asked the question whether WASP-expressing cells might have a selective advantage over WASP-deficient cells. Competitive repopulation experiments were performed in RAG2-deficient mice, since WASP-deficient animals showed a high penetrance of irradiation-induced colitis, preventing their use as recipient mice. Using Southern blot analysis, we documented a selective advantage of wild-type cells in peripheral lymph nodes and spleen, whereas no obvious selective advantage was seen in bone marrow. This finding supports the concept that even the transplantation of a low number of transduced stem cells may lead to therapeutic benefit. In a second series of experiments, we evaluated the feasibility of \*\*\*retroviral\*\*\* gene transfer into WASP-deficient hematopoietic stem cells. Murine scid-hsc from WASP-/- mice were transduced with VSV-G pseudotyped recombinant \*\*\*retroviruses\*\*\* encoding WASP and the marker gene GFP. Upon transplantation of genetically modified stem cells, recipient mice developed a phenotypically normal immune system. Peripheral T-cells showed expression of WASP, as documented by Western blot analysis. Functionally, the specific proliferative defect of WASP-/- T-cells was significantly improved: T-cells from mice transplanted with WASP-deficient stem cells transduced with CMMP-WASP-\*\*\*IRES\*\*\*-GFP showed up to 50% of CD3-induced proliferation levels compared to T-cells from control mice transplanted with wildtype stem cells transduced with the control virus CMMP-GFP. Furthermore, \*\*\*retrovirus\*\*\* -mediated expression of WASP also protected significantly against the onset of colitis in recipient mice. These results provide proof of principle that WASP deficiency can be reversed upon transplantation of \*\*\*retrovirally\*\*\* transduced HSC and may encourage clinical gene therapy trials.

L5 ANSWER 3 OF 324 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.

AN 2001:317213 BIOSIS

DN PREV200100317213

TI Protection of mice from methotrexate and cyclophosphamide induced myelodysplasia by human aldehyde-dehydrogenase and mutated dihydrofolate reductase cDNA gene transfer.

AU Takebe, Naoko (1); Zhao, Shi-Cheng; Banerjee, Debabrata; Bertino, Joseph R.

CS (1) Medicine, University of Maryland, Baltimore, MD USA

SO Blood, ( \*\*\*November 16, 2000\*\*\* ) Vol. 96, No. 11 Part 1, pp. 789a.

print.

Meeting Info.: 42nd Annual Meeting of the American Society of Hematology San Francisco, California, USA December 01-05, 2000 American Society of Hematology

. ISSN: 0006-4971.

DT Conference

LA English

SL English

AB The genetic transfer of drug-resistance to hematopoietic cells is an attractive approach to overcome myelosuppression caused by high dose chemotherapy. Because cyclophosphamide (CTX) and methotrexate (MTX) are commonly used non-cross resistant drugs, generation of dual drug-resistance in hematopoietic cells may allow an increase in dose intensity. We have previously reported in vitro mouse bone marrow progenitor cell protection from 4-hydroxycyclophosphamide (4HC) and methotrexate (MTX) by \*\*\*retroviral\*\*\* gene transfer of human cytosolic class-1 aldehyde-dehydrogenase (ALDH-1) cDNA and a mutated human



dihydrofolate reductase (DHFR; Phe22/Ser31=F/S) gene transfer using SFG based bicistronic MoMLV \*\*\*retroviral\*\*\* vector, SGF-ALDH- \*\*\*IRES\*\*\* -F/S (Takebe N. et al. Blood abstract 554a, 1997). Lethally irradiated mice transplanted with gpAM12-SFG-ALDH- \*\*\*IRES\*\*\* -F/S or mock transduced bone marrow cells were treated with high dose pulse cyclophosphamide (CTX), 200mg/kg daily X 3 or high dose CTX/MTX, 150 mg/kg and 300mg/kg weekly X 2. Animals receiving mock transduced marrow died from CTX and MTX toxicity, whereas mice transplanted with ALDH-1 and mutated DHFR cDNA containing marrow were able to tolerate pulse CTX or weekly CTX/MTX treatment post-transplant. Mice transplanted with transduced marrow and treated with high dose MTX/CTX or high dose CTX alone showed peripheral blood count recovery and maintained their weight, while control mice did not show any blood count recovery and developed weight loss. Genomic DNA from day 11 CFU-S and bone marrow showed evidence of human ALDH-1 cDNA integration by PCR. These data indicate that overexpression of ALDH-1 and mutated DHFR sufficiently induced both 4HC/CTX and MTX resistance in the in vivo mouse model and points to the potential usefulness of this construct to protect patients, requiring high dose CTX and MTX, from myelosuppression.

L5 ANSWER 4 OF 324 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.  
AN 2001:313997 BIOSIS  
DN PREV200100313997

TI Sustained and high level transgene expression in human hematopoietic stem cells transduced by an MSCV/HIV hybrid lentiviral vector.

AU Gao, Zhigang (1); Golob, Jonathan (1); Hawley, Robert G.; Tanavde, Vivek M. (1); Chin, Curt I. (1); Cheng, Linzhao (1)

CS (1) Johns Hopkins Oncology Center, Baltimore, MD USA

SO Blood, ( \*\*\*November 16, 2000\*\*\* ) Vol. 98, No. 11 Part 1, pp. 429a.

print.  
Meeting Info.: 42nd Annual Meeting of the American Society of Hematology San Francisco, California, USA December 01-05, 2000 American Society of Hematology

. ISSN: 0006-4971.

DT Conference

LA English

SL English

AB Oncoretroviral vectors have been used extensively in attempts to transduce human hematopoietic stem-progenitor cells (HSC). We and others have reported that murine stem cell virus (MSCV)-based oncoretroviral vectors transduced HSC efficiently. Lentiviral vectors based on human HIV-1 have been developed recently to transduce HSC. Although high-titer HIV-1 derived vectors have been produced, concerns exist regarding the stability and level of transgene expression that can be achieved following transduction of human HSC. We therefore constructed novel lentivirus vectors which are based on HIV-1 and MSCV. In this study, we present the results obtained with one of the hybrid vectors, pHR-GIN-MU3, in which the U3 region of the HIV LTR was replaced by the U3 region of the MSCV LTR (MU3). For comparison, we also used an oncoretroviral vector, MGIN, containing an identical reporter gene cassette (GFP- \*\*\*IRES\*\*\* -NeoR) controlled by the MSCV LTR. Both vectors directed efficient transgene expression in various human cell lines, indicating that the MU3 enhancer/promoter functioned in the context of the lentiviral backbone. Human cord blood CD34+ cells that had been cultured for 24 hrs in the presence of thrombopoietin, Kit ligand and Flt-3 ligand were transduced twice over 48 hrs with either vector at similar MOI by the spinoculation procedure. FACS analyses and in vitro CFC assays showed that CD34+ cells were transduced by the oncoretroviral and the lentiviral vector at a similar level. To examine transgene expression in the in vivo progeny of transduced human CD34+ cells, the NOD/SCID-transplant assay was used. We observed that 21-40% of the human cells in the bone marrow of NOD/SCID mice expressed the GFP gene introduced by the pHR-GIN-MU3 vector, up to 4 months after transplant. High level GFP expression was observed in both lymphoid, erythroid and myeloid cells. In contrast, low percentages of engrafted human cells expressed GFP from the MGIN vector, presumably because the transduction conditions used were suboptimal for oncoretroviral vectors. Lentiviral vectors such as this MSCV/HIV hybrid vector provide a basis for the design of improved delivery vehicles for gene transfer and expression in human HSC.

L5 ANSWER 5 OF 324 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.  
AN 2001:313287 BIOSIS  
DN PREV200100313287

TI \*\*\*Retrovirus\*\*\* -mediated expression of the base excision repair protein, FPG, protects hematopoietic cells from thiopeta-induced toxicity in vivo.

AU Kobune, M. (1); Xu, Y. (1); Baum, C.; Kelley, M. R. (1); Williams, D. A.

CS (1) Pediatrics, Indiana University School of Medicine, Indianapolis, IN USA

SO Blood, ( \*\*\*November 16, 2000\*\*\* ) Vol. 98, No. 11 Part 1, pp. 461a.

print.  
Meeting Info.: 42nd Annual Meeting of the American Society of Hematology San Francisco, California, USA December 01-05, 2000 American Society of Hematology

. ISSN: 0006-4971.

DT Conference

LA English

SL English

AB Modulation of DNA damage repair activity could lead to new approaches to reduce cytotoxic side effects of chemotherapy. ThioTEPA (TT) induces the formation of N7-aminoethyl (AE) guanine and N1-AE adenine adducts resulting in imidazole ring opening (formamidopyrimidine; Fapy) and is associated with significant myelosuppression in dose-intensive therapies. In E. coli, Fapy lesions are repaired by the Fapy-DNA glycosylase (Fpg) protein. We hypothesized that expression of the Fpg could increase resistance of hematopoietic cells to TT. Fpg-transduced K562 cells demonstrated increased 8-oxodG DNA glycosylase activity and transduced BM progenitor cells were resistant in vitro to TT at multiple concentrations. A \*\*\*retrovirus\*\*\* containing Fpg- \*\*\*IRES\*\*\* -EGFP in a pSF81 backbone (Fpg group; FPG) or the backbone containing \*\*\*IRES\*\*\* -EGFP alone (control group; CN) was used to transduce BM for in vivo analysis. Transduction efficiency was 30-50% and 4-7% (CN and FPG, respectively). Transduced BM cells were transplanted into C57B1/6J mice that were then treated with cycles of 10 mg/kg of TT given (X2) one week apart. In spite of the low transduction efficiency of the FPG vector, peripheral blood leukocytes, hemoglobin and platelet counts of TT-treated FPG mice were significantly higher than CN mice. Platelet counts were well-protected at nadir (238+/-142 X 103/mul vs 100+/-113 X 103/mul, FPG vs CN, p<0.05). After 2 cycles of TT treatment, the numbers of EGFP+ progenitor cells in BM of treated FPG mice were significantly higher compared with those of CN mice, in spite of the initial transduction efficiency (387+/-88.9 vs 112+/-121 X 105 cells, p<0.05). Splenic and thymic cellularity of FPG mice

were also significantly higher (spleen 88.9+/-18.9 X 106 vs 31.1+/-8.5 X 106, p<0.01; thymus 9.5+/-5.9 X 105 vs 1.5+/- 1.0 X 106, FPG vs CN, respectively, p<0.05). Selective pressure was also demonstrated by an increase in the proportion of EGFP "bright" cells after TT. Mean fluorescence intensity (MFI) of peripheral mononuclear cells (PBMC) of FPG group was increased after 1 cycle of TT treatment compared with pretreatment MFI (1052+/-747 vs 523+/-205, p<0.05), while the MFI of CN and non-treated FPG mice were not changed. These results show that expression of the Fpg protein protects hematopoietic cells from TT-induced DNA damage and BM cells highly expressing bacterial Fpg selectively survive during TT in vivo treatment. Fpg may provide a novel approach to preventing TT-induced toxicity of primary hematopoietic cells.

L5 ANSWER 6 OF 324 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.

AN 2001:311927 BIOSIS

DN PREV200100311927

TI Myeloma cells homing to the bone marrow is directed by CXCR4/SDF-1 interactions.

AU Woodruff, Jeffrey E. (1); Engel, Barbara C.; Epstein, Joshua (1)

CS (1) Myeloma and Transplantation Research Center, University of Arkansas for Medical Sciences, Little Rock, AR USA

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. ISSN: 0006-4971.

DT Conference

LA English

SL English

AB Multiple Myeloma is characterized by malignant plasma cell infiltration throughout the bone marrow, resulting in lytic bone lesions, a devastating manifestation of this disease. The mechanism controlling myeloma cells homing to the bone marrow have not been elucidated. We have previously reported that primary myeloma cells express the chemokine receptor CXCR4 and migrate in vitro in response to its ligand SDF-1. To further determine whether this mechanism is responsible for active myeloma cell homing to the bone marrow in vivo, we investigated the dissemination of myeloma cells engineered to differentially express CXCR4 in SCID mice. ARP-1 cells, a cell line established from the bone marrow of a myeloma patient, express low levels of CXCR4. ARP-1 cells were transduced with CXCR4 to generate stable transfectants (ARP-1X), with constitutive expression 20 fold higher than that of parental cells. To reduce endogenous expression and to minimize the effect of in vivo induction of CXCR4 expression seen in preliminary experiments, ARP-1 cells were also transduced with the \*\*\*retroviral\*\*\* SDF-1 intralidase vector MND-SDF-KDEL. \*\*\*IRES\*\*\* -eGFP (MSKIE). SCID mice were inoculated intravenously with ARP-1X or ARP-1neo/MSKIE cells. When tumor developed, the presence of tumor cells in the different organs was determined using CD38/CD45 (for ARP-1X) or GFP flow cytometry (for ARP-1neo/MSKIE). All mice injected with ARP-1X cells had tumor cells in their femurs (11.9%+/-2.1) and in their vertebrae (25.3%+/-28.7). In contrast, only two of the 5 mice injected with ARP-1neo/MSKIE cells had tumor cells in the femur (0.08% and 0.01%), one of these also had 0.1% tumor cells in its vertebrae. The difference between the two groups in bone marrow plasmacytosis was statistically significant (p=0.02). These results demonstrate unequivocally that CXCR4 directs active migration of myeloma cell towards the bone marrow.

L5 ANSWER 7 OF 324 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.

AN 2001:311903 BIOSIS

DN PREV200100311903

TI BCR-ABL induces normal erythropoiesis in the absence of JAK2.

AU Ghaffari, Sagh (1); Kitidis, Claire (1); Neubauer, Hans; Pfeiffer, Klaus; Lodish, Harvey (1)

CS (1) Whitehead Institute, Cambridge, MA USA

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DT Article, Conference

LA English

SL English

AB We have shown previously that the constitutively active tyrosine kinase BCR-ABL oncoprotein (P210) induces red cell formation in Epor-/- fetal liver cells (FLC). JAK2 is an integral component of Epor where it initiates the stimulation of downstream signaling pathways. JAK2 function is crucial for definitive erythropoiesis, as JAK2-deficient mice die from fetal anemia by embryonic day 12 or 13, similar to Epor-/- mice; however, JAK2-/- embryos suffer from a more severe defect. We have found JAK2 to be constitutively phosphorylated in the erythroleukemic HCD57 cell line expressing P210.JAK2 phosphorylation is increased significantly upon Epo stimulation in HCDP210 cells as compared to the parental HCD57 cells. We find JAK2 to be also constitutively phosphorylated in primary FLC \*\*\*retrovirally\*\*\* expressing BCR-ABL (P210). We sought to determine whether JAK2 is required for red cell formation by P210. Using a bicistronic MSCV- \*\*\*IRES\*\*\* -GFP vector, we generated high titer \*\*\*retroviral\*\*\* supernatants to transduce day12 JAK2-/- FLC to express either P210, JAK2, or an empty vector. The cells were cultured in the presence of Steel factor (SF) and IL-6, and two days post-infection, GFP+ FLC were analyzed for their frequency of cells expressing the red cell marker Ter119. GFP+ JAK2-/- FLC infected with P210 generated as many Ter119+ cells as the ones infected with JAK2 and cultured in the presence of Epo+SF+IL-6. In addition, GFP+Ter119- FLC were selectively sorted and plated in methylcellulose in the presence of either SF+IL-6 (P210-infected cells) or SF+IL-6+Epo (JAK2 or control vector) and BFU-E colonies counted after 9 days. In the absence of Epo signaling, P210-infected cells generated as many BFU-E colonies as the ones infected with JAK2 and cultured in the presence of Epo. However, the presence of at least SF was required for P210-infected cells to generate BFU-E colonies from JAK2-/- FLC, an effect distinct from the one seen with Epor-/- FLC.P210 or JAK2-generated BFU-E appeared to have similar size and morphology. RT-PCR on individual BFU-Es detected the expression of GATA-1, globin gene and Epor in both P210 and JAK2-generated colonies. GFP+Ter119- cells were also assayed for their CFU-E content. P210-infected cells cultured in the presence of SF generated as many benzidine positive CFU-E as JAK2-infected cells cultured in SF+Epo. Taken together these data indicate that although JAK2 is constitutively phosphorylated in BCR-ABL-expressing erythroid cells, BCR-ABL does not require JAK2 for inducing red cell formation.

L5 ANSWER 8 OF 324 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.

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DT Article

LA English

AB Most eukaryotic mRNAs are translated by a "scanning ribosome" mechanism.

We have found that unlike the type member of the genus Tobamovirus, translation of the 3'-proximal \*\*\*coat\*\*\* \*\*\*protein\*\*\* ( \*\*\*CP\*\*\* ) gene of a crucifer infecting tobamovirus (crTMV) (Dorokhov et al., 1993; 1994) occurred in vitro by an internal ribosome entry mechanism. Three types of synthetic dicistronic RNA transcripts were constructed and translated in vitro: (i) "MP- \*\*\*CP\*\*\* -3'NTR" transcripts contained movement protein (MP) gene, \*\*\*CP\*\*\* gene and the 3'-nontranslated region of crTMV RNA. These constructs were structurally equivalent to dicistronic subgenomic RNAs produced by tobamoviruses in vivo. (ii) "DELTA-NPT- \*\*\*CP\*\*\* " transcripts contained partially truncated neomycin phosphotransferase I gene and \*\*\*CP\*\*\* gene. (iii) " \*\*\*CP\*\*\* -GUS" transcripts contained the first \*\*\*CP\*\*\* gene and the gene of Escherichia coli beta-glucuronidase (GUS) at the 3'-proximal position. The results indicated that the 148-nt region upstream of the \*\*\*CP\*\*\* gene of crTMV RNA contained an internal ribosome entry site (IREScp) promoting internal initiation of translation in vitro. Dicistronic \*\*\*IRES\*\*\* - \*\*\*CP\*\*\* , containing chimeric mRNAs with the 5'-terminal stem-loop structure preventing translation of the first gene (MP, DELTA-NPT, or \*\*\*CP\*\*\* ), expressed the \*\*\*CP\*\*\* or GUS genes despite their 3'-proximal localization. The capacity of crTMV IREScp for mediating internal translation distinguishes this \*\*\*CP\*\*\* tobamovirus from the well-known type member of the genus, TMV U1. The equivalent 148-nt sequence from TMV RNA was incapable of mediating internal translation. Two mutants were used to study structural elements of IREScp. It was concluded that integrity of \*\*\*IRES\*\*\* - \*\*\*CP\*\*\* was essential for internal initiation. The crTMV provides a new example of internal initiation of translation, which is markedly distinct from IRESs shown for picomaviruses and other viral and eukaryotic mRNAs.

L4 ANSWER 12 OF 12 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.DUPLICATE

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AN 1995:361093 BIOSIS

DN PREV199598375393

TI MRNAs containing the unstructured 5' leader sequence of alfalfa mosaic virus RNA 4 translate inefficiently in lysates from poliovirus-infected HeLa cells.

AU Hann, Louane E.; Gehrke, Lee (1)

CS (1) M.I.T. Build. E25-545, Cambridge, MA 02139 USA

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ISSN: 0022-538X.

DT Article

LA English

AB Poliovirus infection is accompanied by translational control that precludes translation of 5'-capped mRNAs and facilitates translation of the uncapped poliovirus RNA by an internal initiation mechanism. Previous reports have suggested that the capped alfalfa mosaic virus \*\*\*coat\*\*\* \*\*\*protein\*\*\* mRNA (AIMV \*\*\*CP\*\*\* RNA), which contains an unstructured 5' leader sequence, is unusual in being functionally active in extracts prepared from poliovirus-infected HeLa cells (PI-extracts). To identify the cis-acting nucleotide elements permitting selective AIMV \*\*\*CP\*\*\* expression, we tested capped mRNAs containing structured or unstructured 5' leader sequences in addition to an mRNA containing the poliovirus internal ribosome entry site ( \*\*\*IRES\*\*\* ). Translations were performed with PI-extracts and extracts prepared from mock-infected HeLa cells (MI-extracts). A number of control criteria demonstrated that the HeLa cells were infected by poliovirus and that the extracts were translationally active. The data strongly indicate that translation of RNAs lacking an internal ribosome entry site, including AIMV \*\*\*CP\*\*\* RNA, was severely compromised in PI-extracts, and we find no evidence that the unstructured AIMV \*\*\*CP\*\*\* RNA 5' leader sequence acts in cis to bypass the poliovirus translational control. Nevertheless, cotranslation assays in the MI-extracts demonstrate that mRNAs containing the unstructured AIMV \*\*\*CP\*\*\* RNA 5' untranslated region have a competitive advantage over those containing the rabbit alpha-globin 5' leader. Previous reports of AIMV \*\*\*CP\*\*\* RNA translation in PI-extracts likely describe inefficient expression that can be explained by residual cap-dependent initiation events, where AIMV \*\*\*CP\*\*\* RNA translation is competitive because of a diminished quantitative requirement for initiation factors.

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